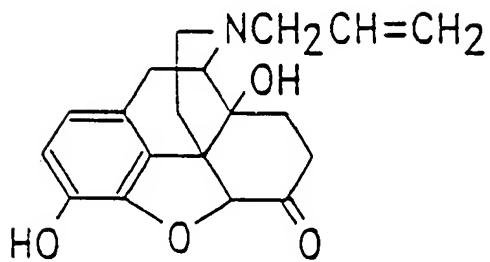
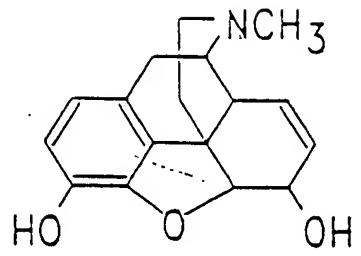
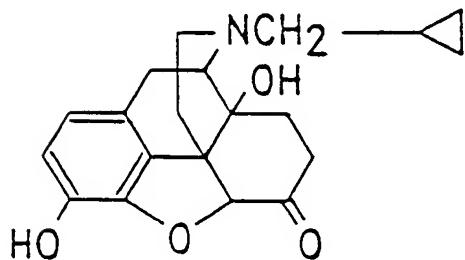


APPROVED	O.G. FIG.
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Morphine



Naloxone



Naltrexone (R=O)
Nalmefene (R=CH₂)

FIG. 1

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1. BSS
2. Et 1mM
3. Et 1μM
4. Et 1nM
5. Et 1uM
6. BSS

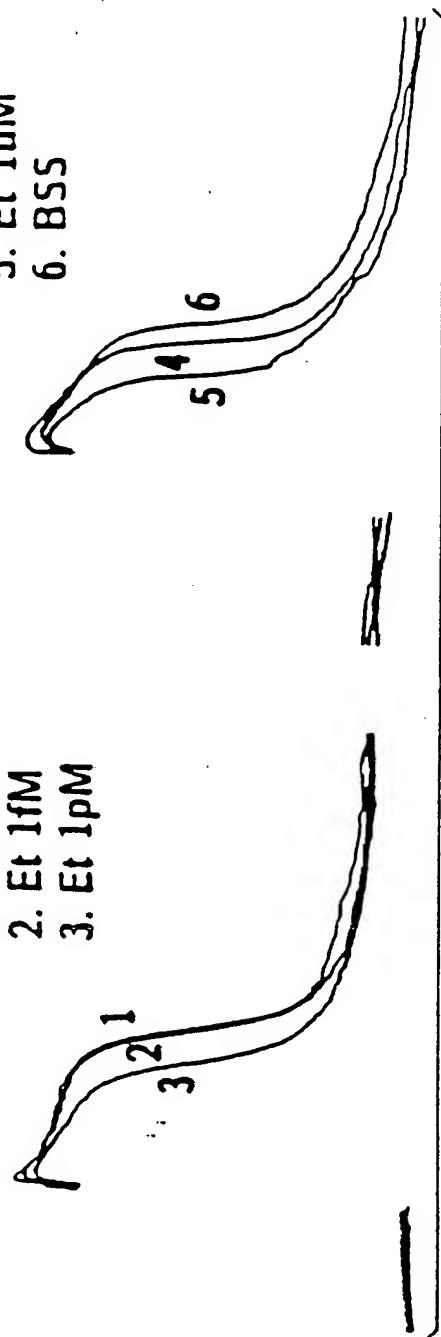


FIG. 2A

1. BSS
2. Et 1mM
3. NLX 10nM
4. NLX + Et 1nM
5. NLX + Et 1uM

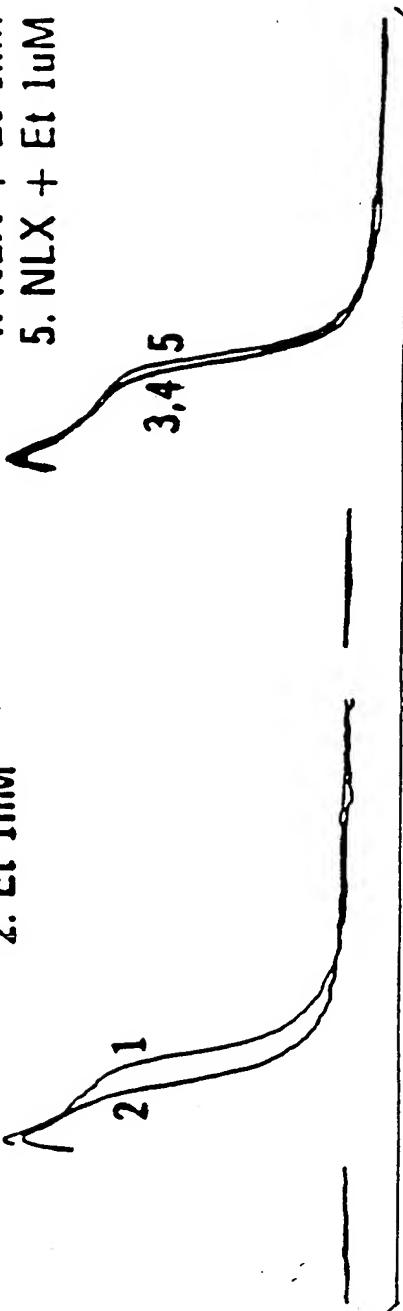


FIG. 2B

APPROVED	O.G. FIG.
BY	CLASS SUBCLASS
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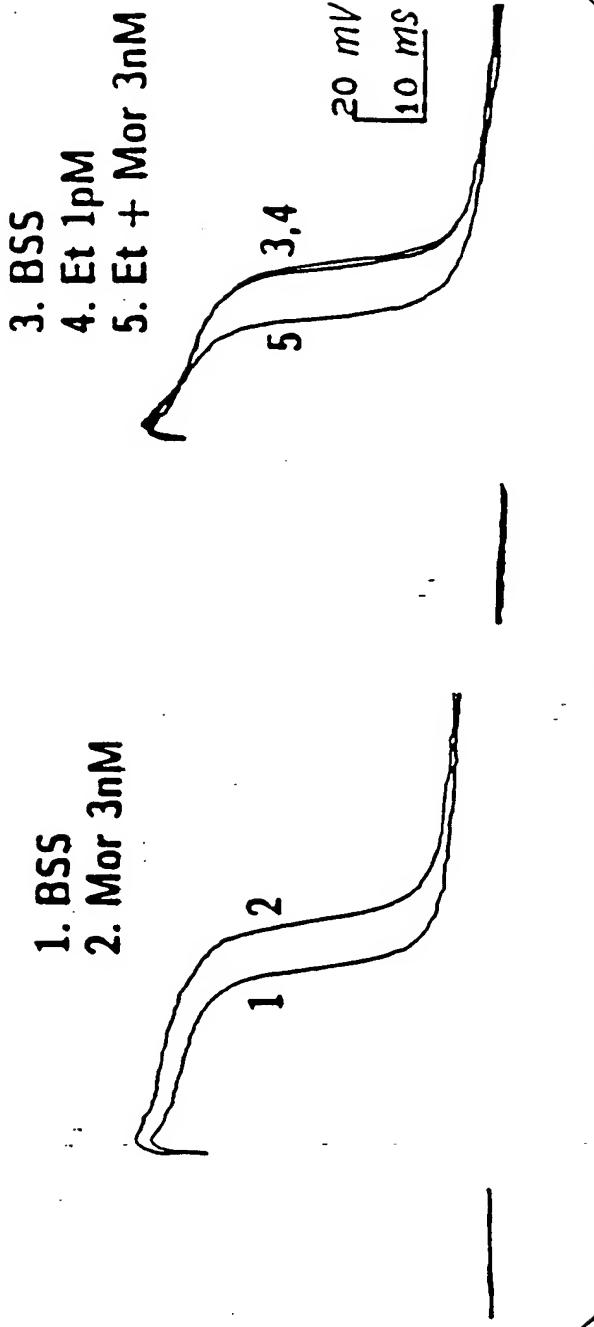


FIG. 2C

APPROVED	O.G. FIG.	—
BY	CLASS	SUBCLASS
DRAFTSMAN	514	285

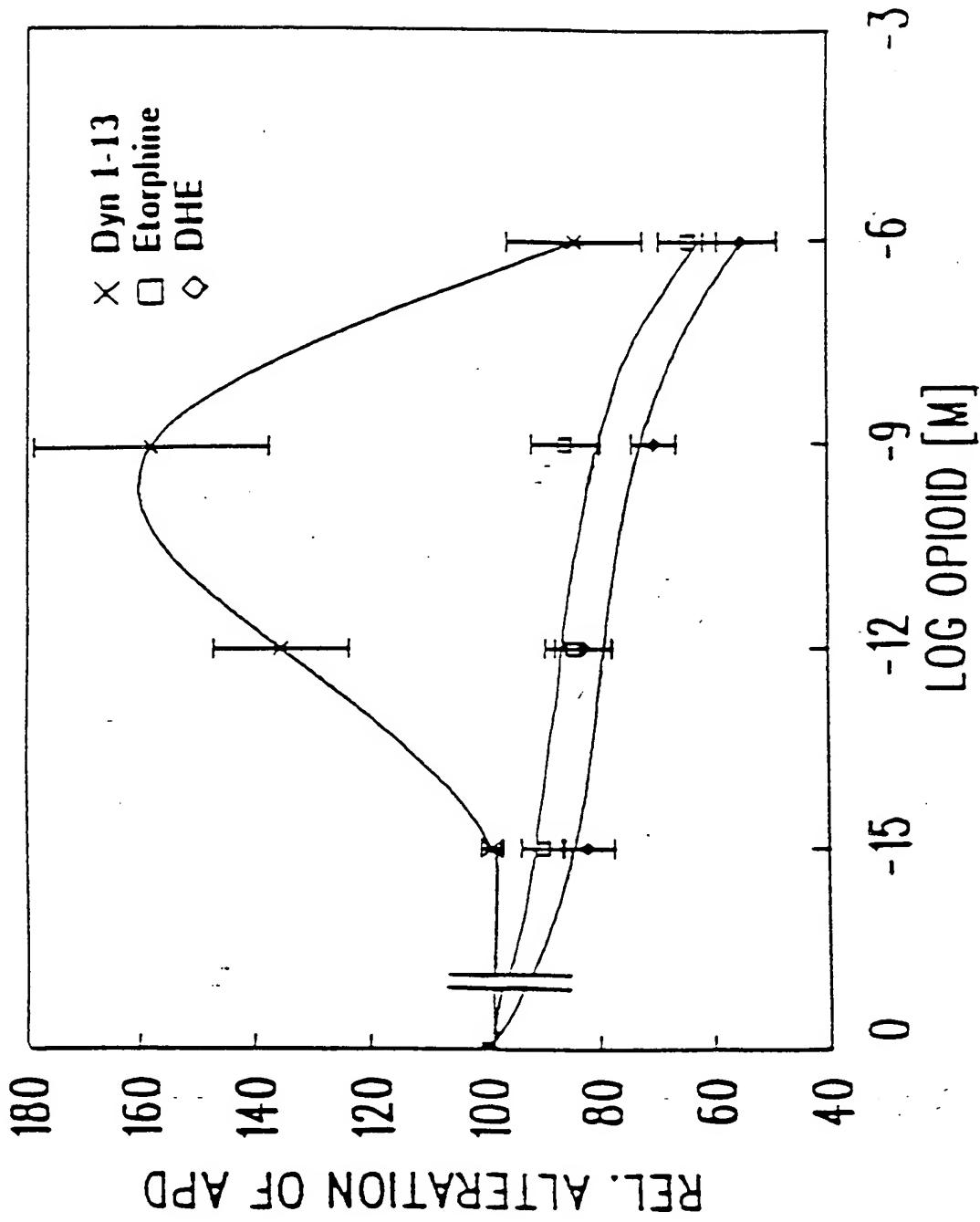


FIG. 3

APPROVED BY	O. G. FIG.	
DRAFTSMAN	514	CLASS SUBCLASS
		285

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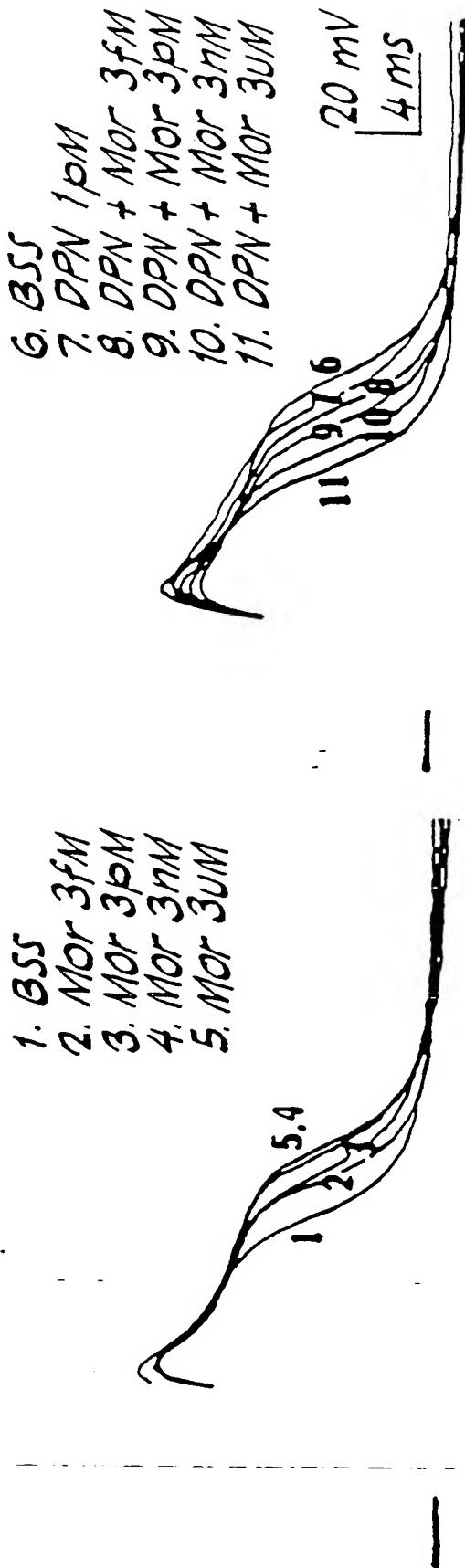


FIG. 4A

APPROVED	O.C. FIG.	—
BY	CLASS	SUBCLASS
DRAFTSMAN	714	285

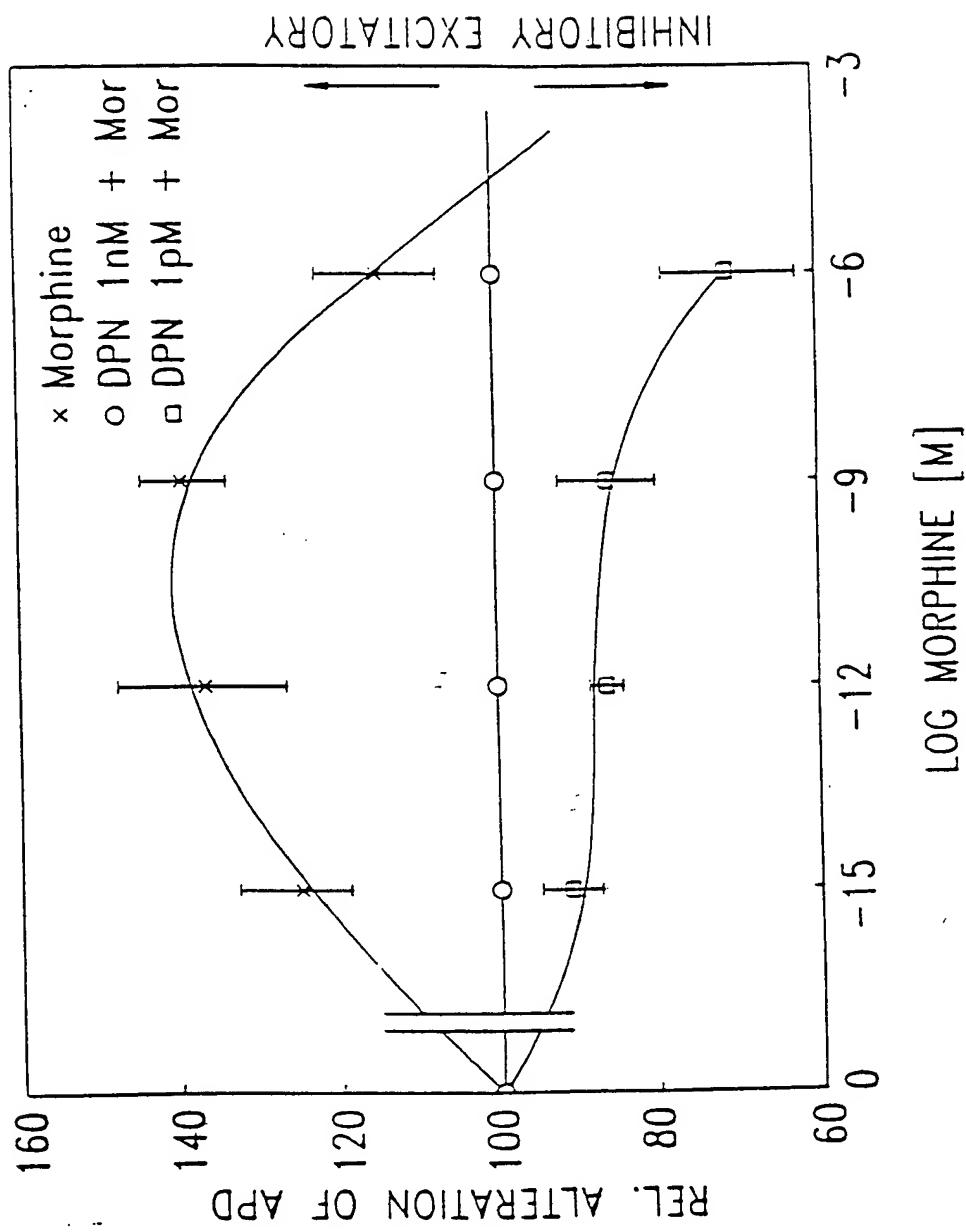


FIG. 4B.

APPROVED	O.G. FIG.	—
BY	CLASS	SUBCLASS
DRAFTSMAN	514	285

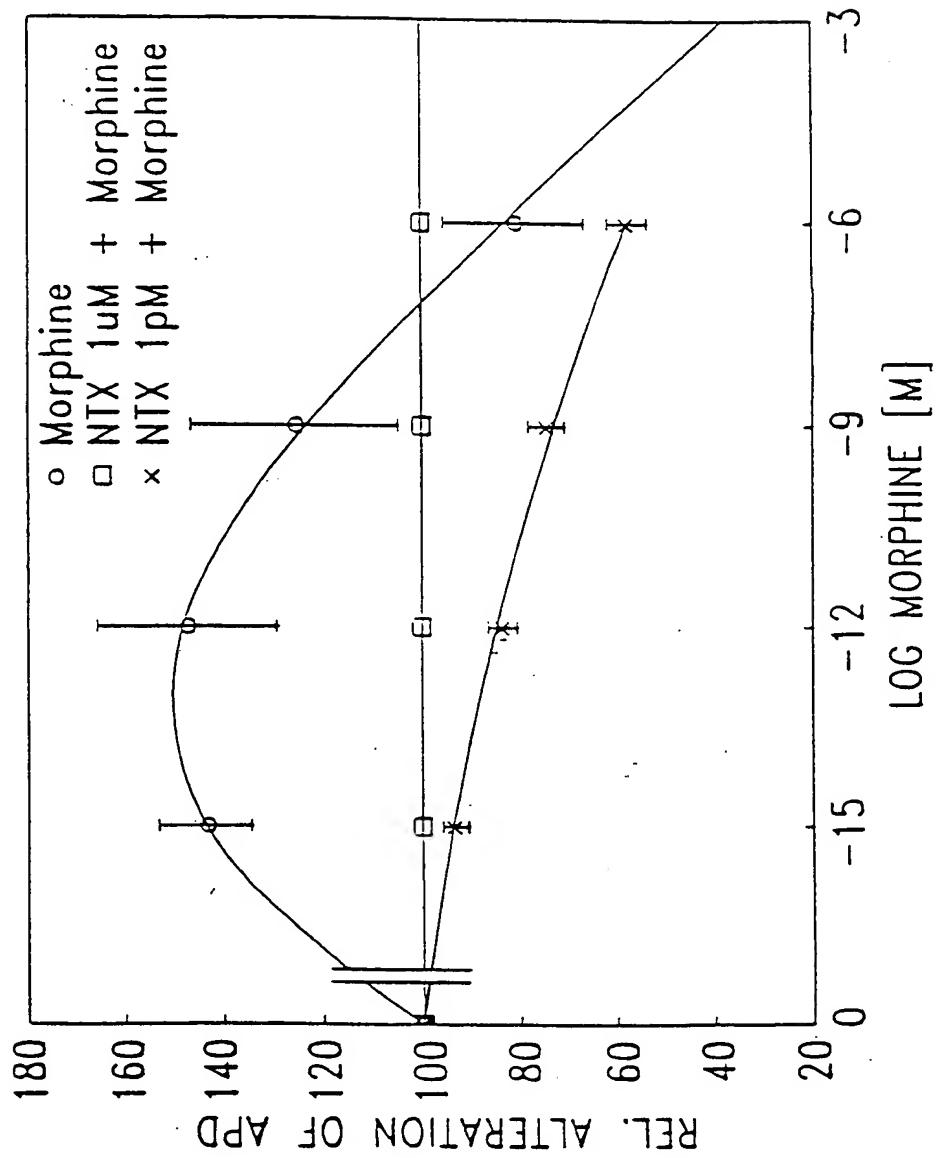


FIG. 5

APPROVED	O.G. FIG.
BY	
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CLASS	SUBCLASS

Alteration of Action Potential Duration (APD)			
Acute Test	(APD shortening: ↓ ; APD prolongation: ↑ ; No APD change: 0)		Chronic Co-treatment with Mor + Antag. at Excit. Op. Rec. (pM)
	Naive DRG Neurons		
	Control BSS	BSS + Antag. at Excit.Op.Rec. (pM)	After Washout with BSS
1 - 10 μ M morphine	↓ (inhibitory) ("analgesia")	↑ ("tolerance")	↑
pM - nM morphine	↓ ("excitatory" "antianalgesia")	↑ (unmasking of inhibitory effects)	↑
\sim fM morphine or dyn A-(1-13)	0	0 (excitatory supersensitivity)	0
nM naloxone	0	0 ("dependence") ("withdrawal effect")	0

FIG. 6

APPROVED BY	O.G. FIG.
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	CLASS SUBCLASS 285

CONFIDENTIAL

Co-Treatment with Low-dose Naltrexone Enhances Morphine's Antinociceptive Potency

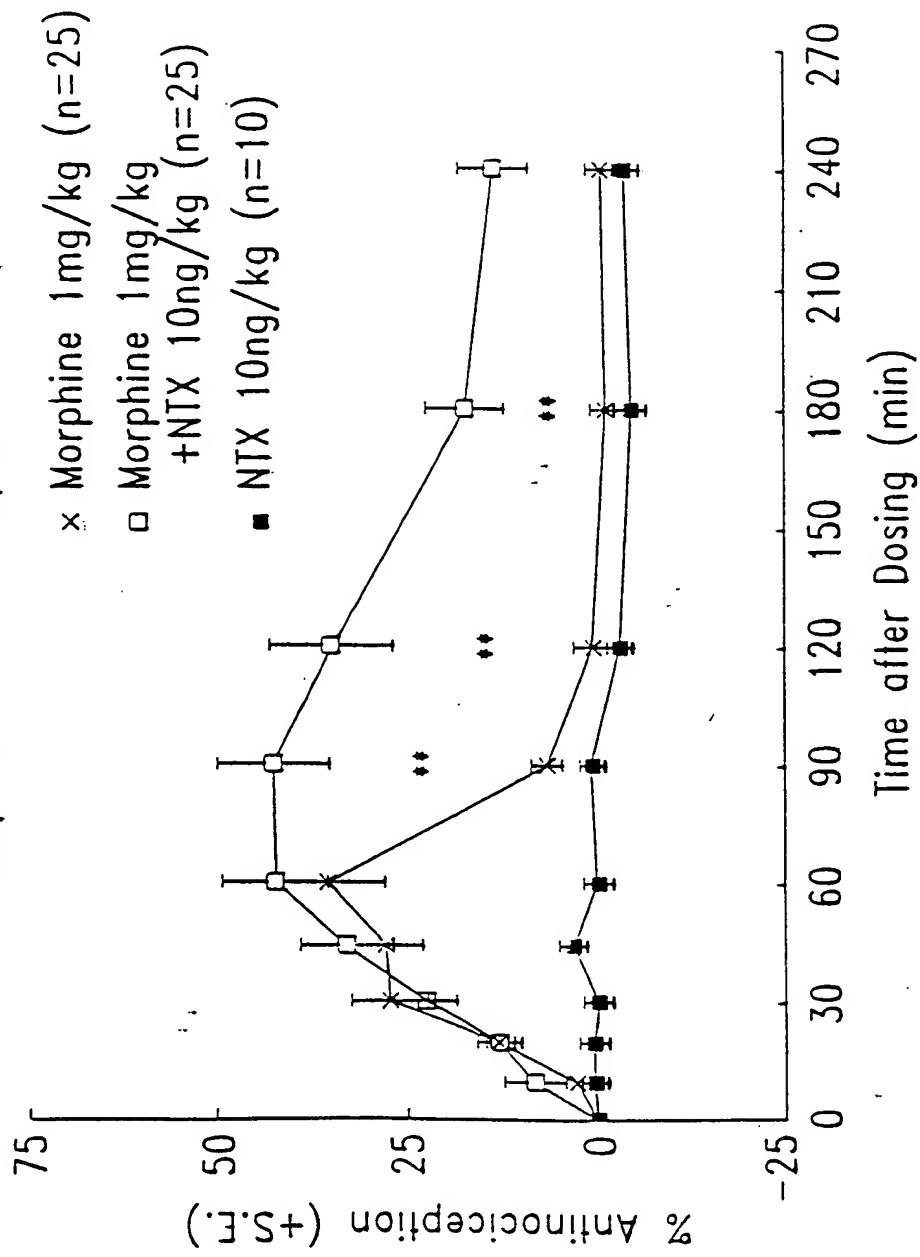


FIG. 7

APPROVED	O.G. FIG.
BY	
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CLASS	SUBCLASS

SEARCHED INDEXED

Co-Treatment with Low-Dose Naltrexone Attenuates
Acute and Chronic Morphine Dependence

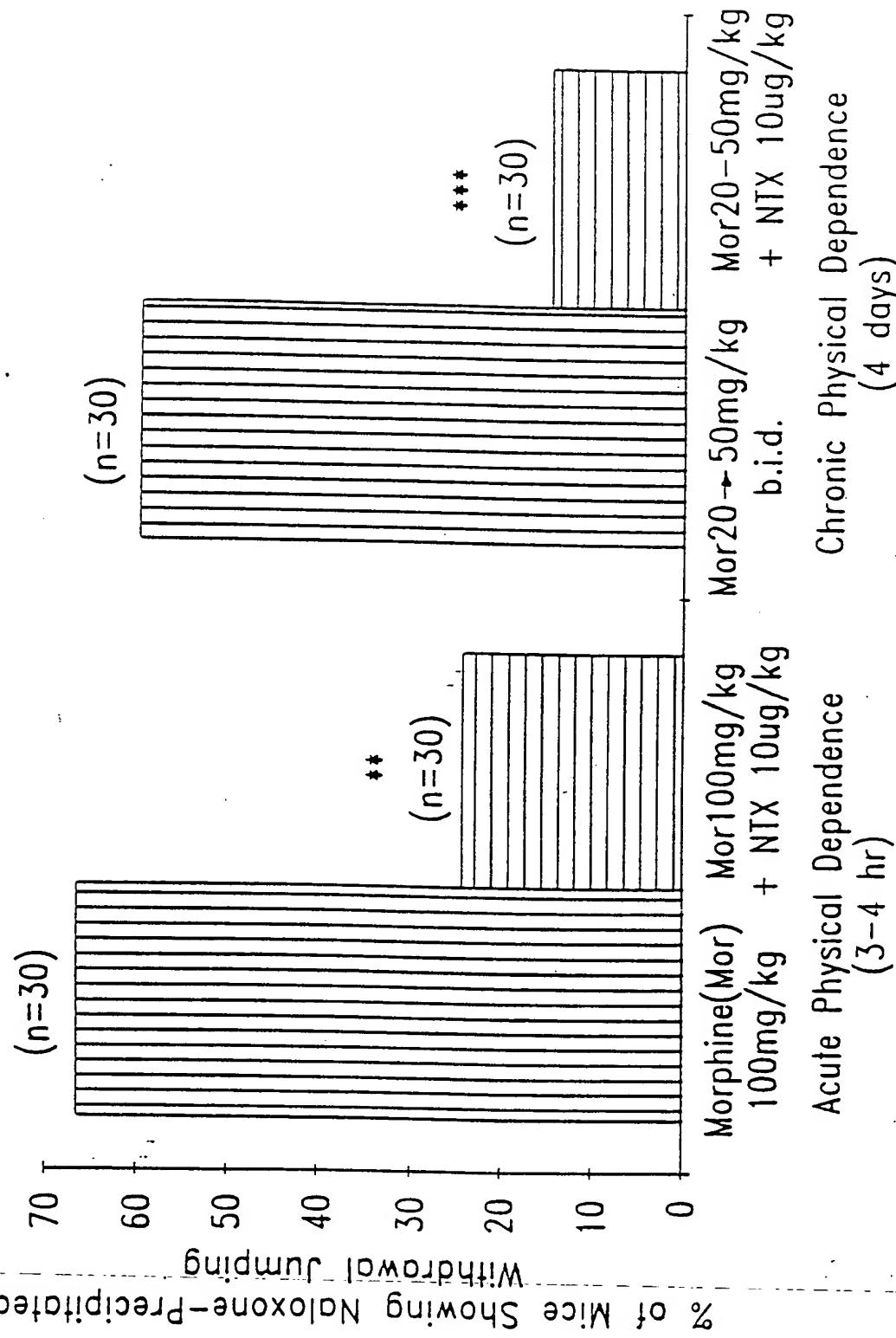


FIG. 8

APPROVED BY	O.G. FIG. —
DRAFTSMAN	574
	28.5

CONFIDENTIAL

Co-Treatment with Ultra-Low Dose Naloxone Enhances
Morphine Antinociceptive Potency

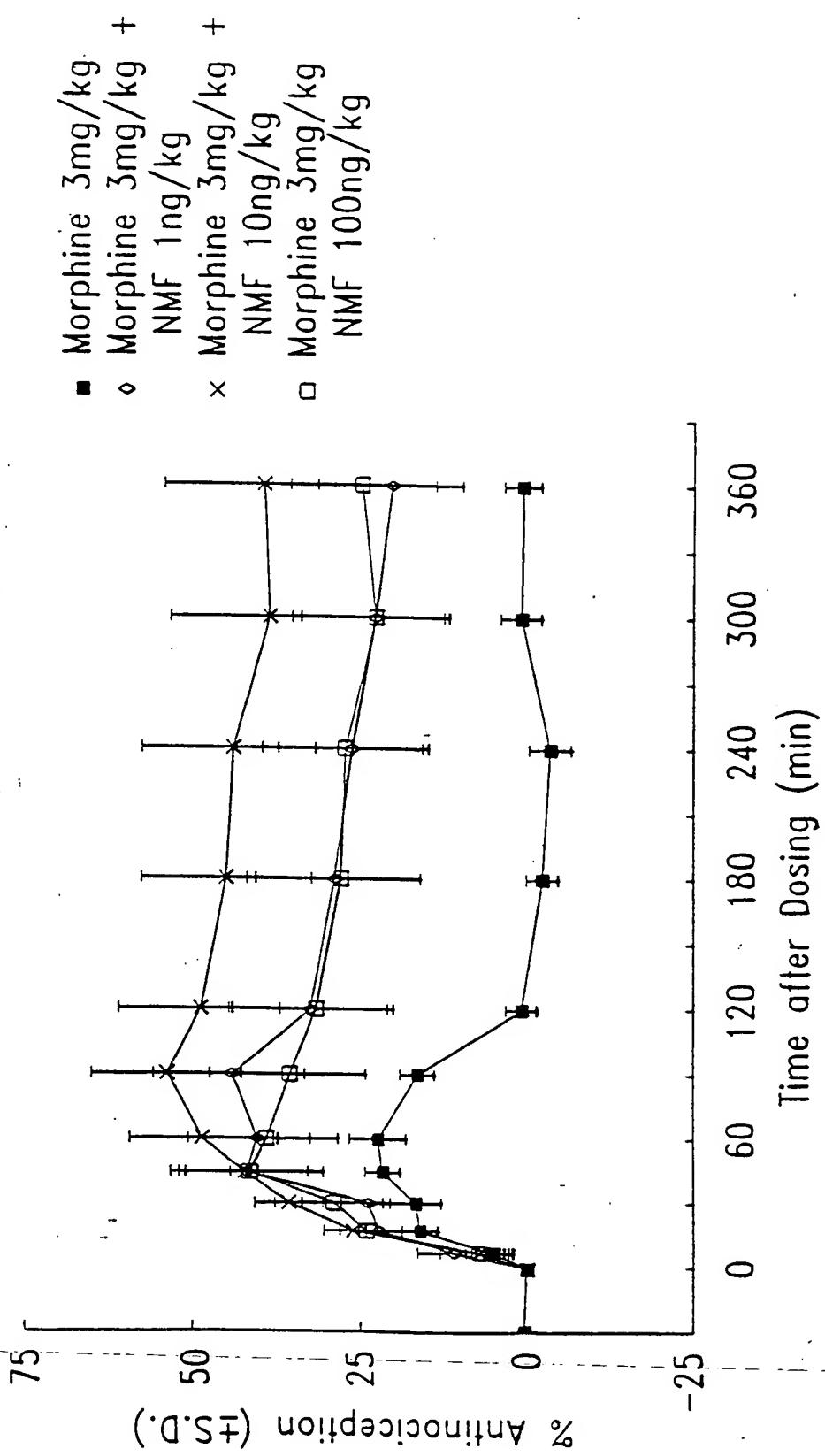


FIG. 9

APPROVED	O.G. FIG.
BY	—
DRAFTSMAN	CLASS SUBCLASS 514 285

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Co-Treatment with Low-Dose Naltrexone or Nalmefene Attenuates Acute Morphine Dependence

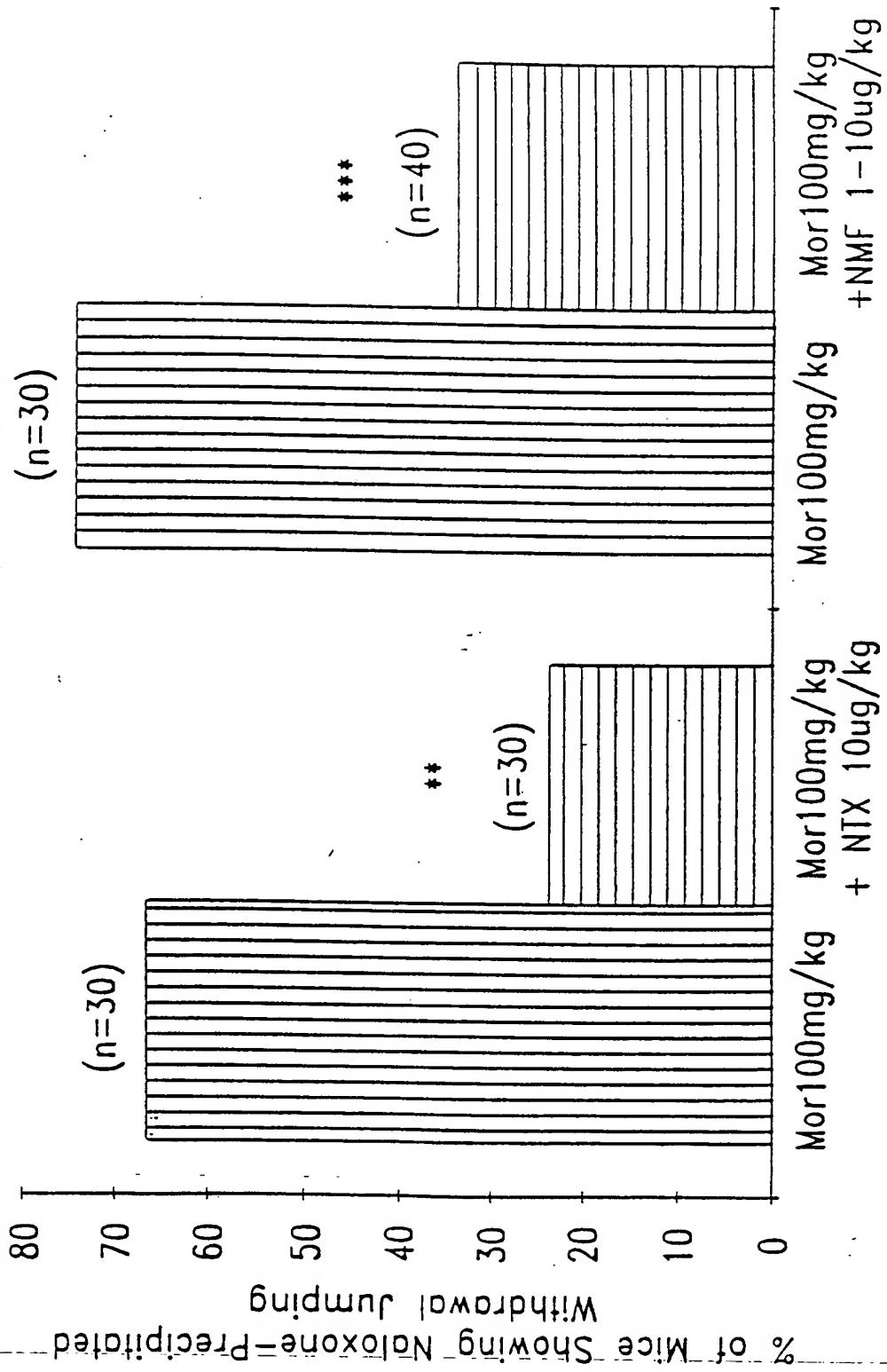


FIG. 10